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Services**Identification of a novel signal transduction surface molecule on human cytotoxic lymphocytes.****Valiante NM, Trinchieri G.**56 Nicholas M
Giorgio

Wistar Institute of Anatomy and Biology, Philadelphia, Pennsylvania 19104.

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In this study, we have used a newly generated monoclonal antibody (mAb C1.7) to identify a novel 38-kD signal-transducing surface molecule (p38) expressed by lymphocyte subsets capable of cell-mediated cytotoxicity. Virtually all CD16+/CD56+ natural killer (NK) cells and approximately half of CD8+ (T cell receptor [TCR] alpha/beta+) T cells and TCR-gamma/delta+ T cells express the p38 surface molecule. Stimulation of p38 on NK cells with mAb C1.7 activated cytotoxicity, induced lymphokine production, and initiated polyphosphoinositol turnover and [Ca²⁺]_i increases. Unlike other NK cell surface molecules that activate cytotoxicity, p38 stimulation did not result in the release of the granule enzyme N-carbobenzoxymethyl-L-thiobenzyl ester-esterase even under conditions in which mAb C1.7 induced NK cell-mediated redirected lysis of Fc gamma R+ target cells. Activated (recombinant interleukin 2 [rIL-2], 5 d) CD8+ T cells mediated non-major histocompatibility complex (MHC)-restricted cytotoxicity, and the CD8+/p38+ subset contained the overwhelming majority of this activity. F(ab')₂ fragments of mAb C1.7 inhibited non-MHC-restricted cytotoxicity mediated by resting NK cells and rIL-2-cultured T cells but did not affect spontaneous cytotoxicity mediated by activated, cultured NK cells. Taken as a whole, our results suggest that p38 may have a direct role in the recognition, signal transduction, and/or lytic mechanisms of non-MHC-restricted cytotoxicity.

PMID: 8376943 [PubMed - indexed for MEDLINE]

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Inventor Name Search Result

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Last Name = TRINCHIERI

First Name = GIORGIO

Application#	Patent#	Status	Date Filed	Title	Inventor Name
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<u>07307817</u>	Not Issued	164	02/07/1989	NATURAL KILLER CELL STIMULATORY FACTOR	TRINCHIERI , GIORGIO
<u>07584941</u>	<u>5457038</u>	150	09/18/1990	NATURAL KILLER STIMULATORY FACTOR	TRINCHIERI , GIORGIO
<u>08229282</u>	Not Issued	168	04/18/1994	COMPOSITIONS AND METHODS FOR USE OF IL-12 AS AN ADJUVANT	TRINCHIERI , GIORGIO
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<u>08307280</u>	<u>5688690</u>	150	09/16/1994	HUMAN CYTOTOXIC LYMPHOCYTE SIGNAL TRANSDUCTION SURFACE PROTEIN (P 38) AND MONOCLONAL ANTIBODIES THERETO	TRINCHIERI , GIORGIO
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- ☐ 4: Peritt D, Sesok-Pizzini DA, Schretzenmair R, Macgregor RR, Valiante NM, Tu X, Trinchieri G, Kamoun M. Related Articles
C1.7 antigen expression on CD8+ T cells is activation dependent: increased proportion of C1.7+CD8+ T cells in HIV-1-infected patients with progressing disease.
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- ☐ 5: Tangye SG, Lazetic S, Woollatt E, Sutherland GR, Lanier LL, Phillips JH. Related Articles, Nucleotide, OMIM, Protein
Cutting edge: human 2B4, an activating NK cell receptor, recruits the protein tyrosine phosphatase SHP-2 and the adaptor signaling protein SAP.
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- ☐ 6: Robertson MJ, Cochran KJ, Cameron C, Le JM, Tantravahi R, Ritz J. Related Articles
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